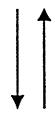
FIG.1

[SEQ. ID NO: 3] X-C-C-T-T-G-A-G-A-T-T-T-C-C-C-T-C

G-G-A-A-C-T-C-T-A-A-A-G-G-G-A-G-X
[SEQ. ID NO: 4]



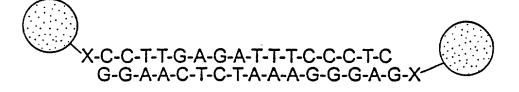


FIG.2

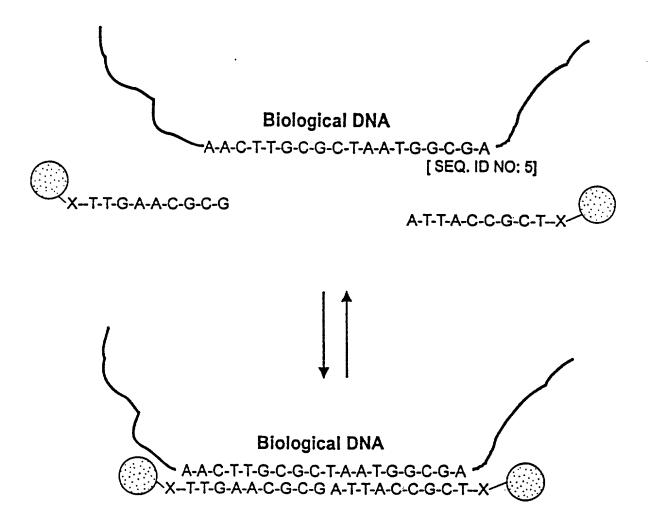
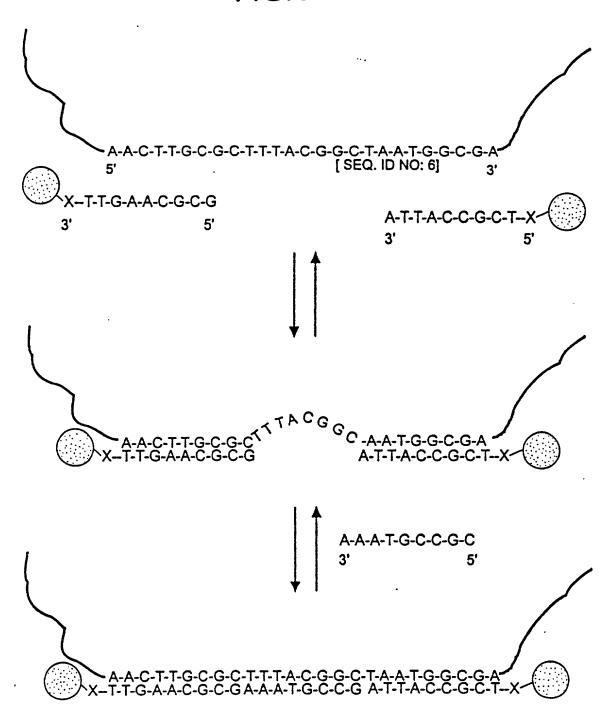
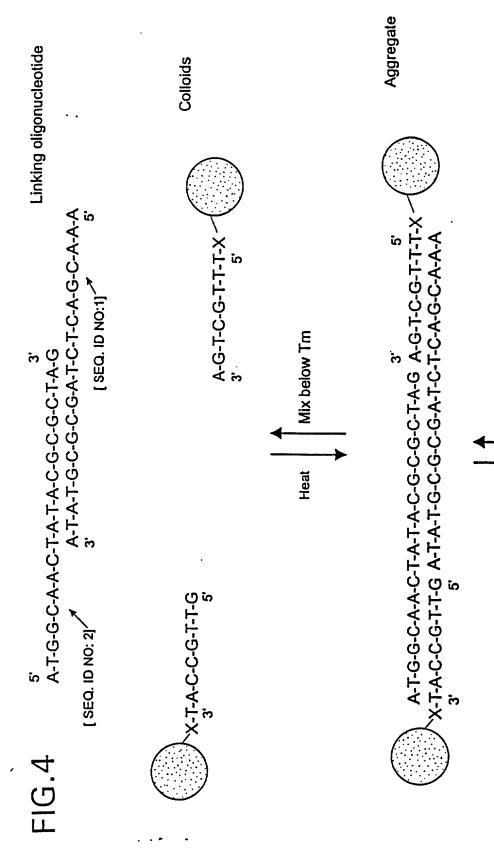


FIG.3





Precipitate (formed by further cross-linking)

Stand below Tm

Heat

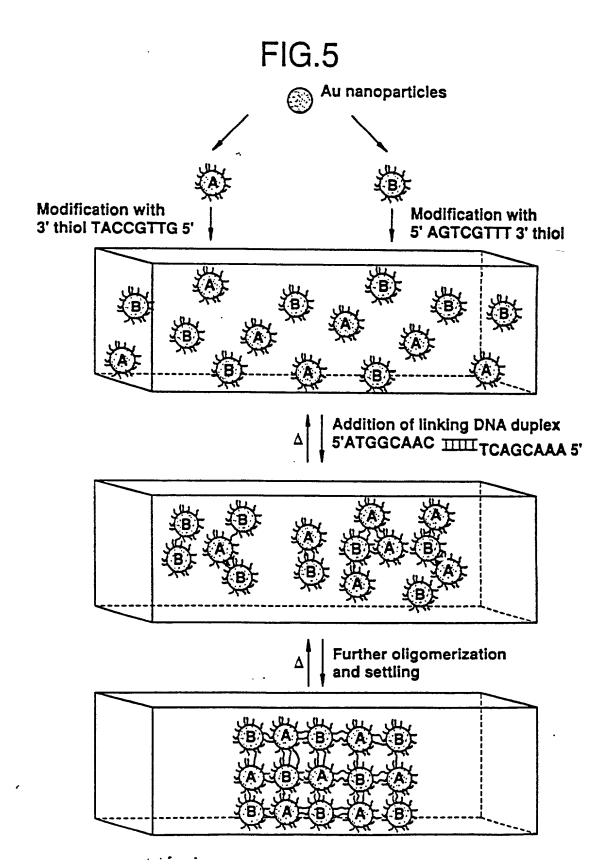
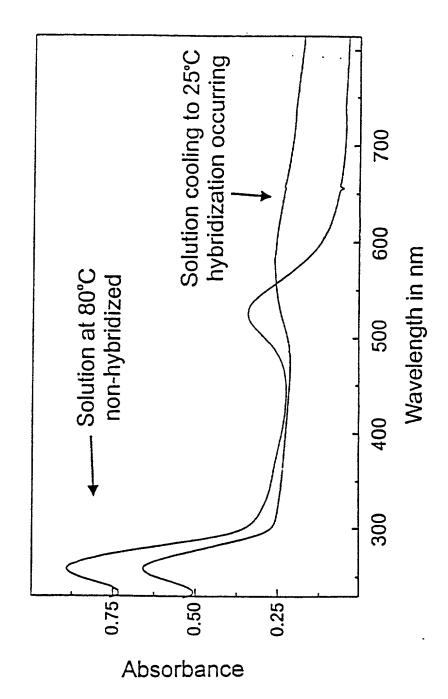
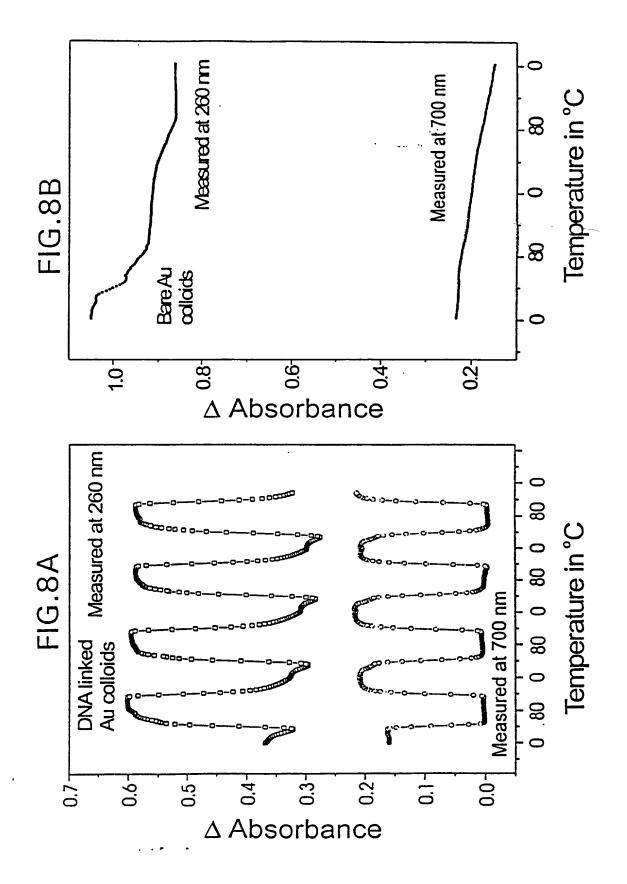




FIG.6A FIG.6B FIG.6C

FIG.7





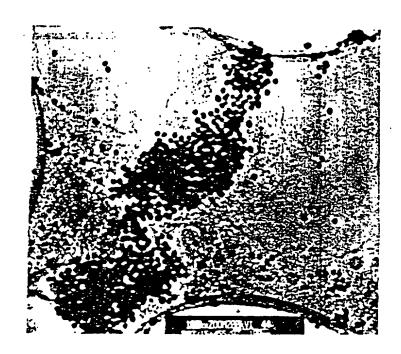


FIG.9A

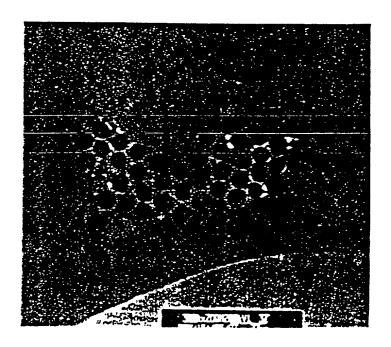
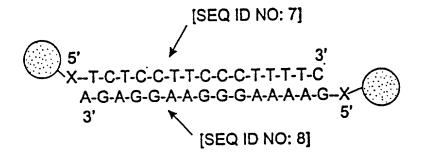
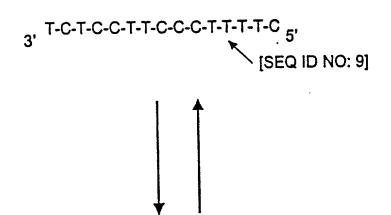


FIG.9B

FIG.10





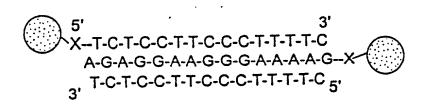
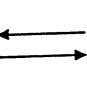


FIG.11

[SEQ. ID NO: 10]
s-A-T-G-G-C-A-A-C-T-A-C-G-C-G-C-T-A-G-A-G-T-C-G-T-T-T
5'

T-A-C-C-G-T-T-G-A-T-A-T-G-C-G-C-G-A-T-C-T-C-A-G-C-A-A-A-S-C 5'
3'
[SEQ. ID NO: 11]



T-A-C-C-G-T-T-G-A-T-A-T-G-C-G-C-G-A-T-C-T-C-A-G-C-A-A-A s-A-T-G-G-C-A-A-C-T-A-T-A-C-G-C-G-C-T-A-G-A-G-T-C-G-T-T-T

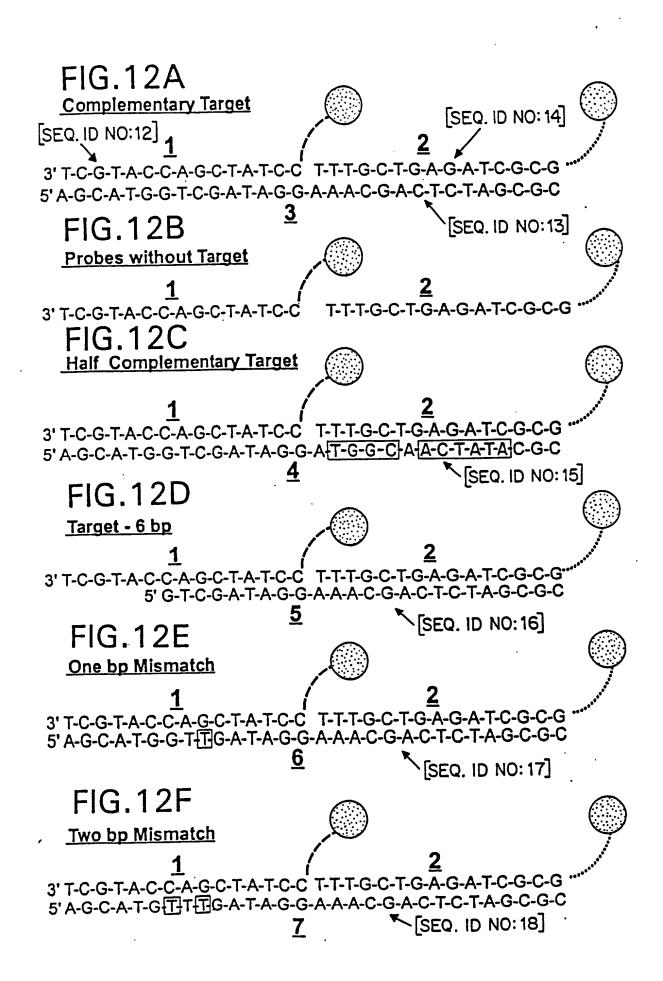
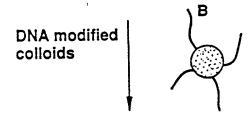


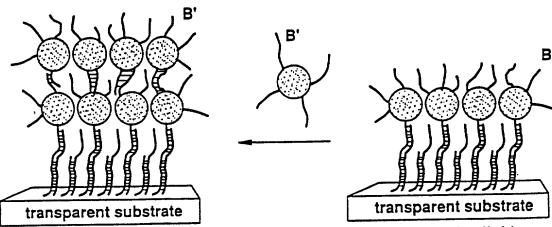
FIG.13A

transparent substrate

Modified DNA chemisorbed onto solid substrate

Analyte DNA hybridized onto substrate





Dark areas where nanoparticle aggregates are linked to substrate surface by analyte DNA

DNA modified colloids hybridized to bound analyte DNA

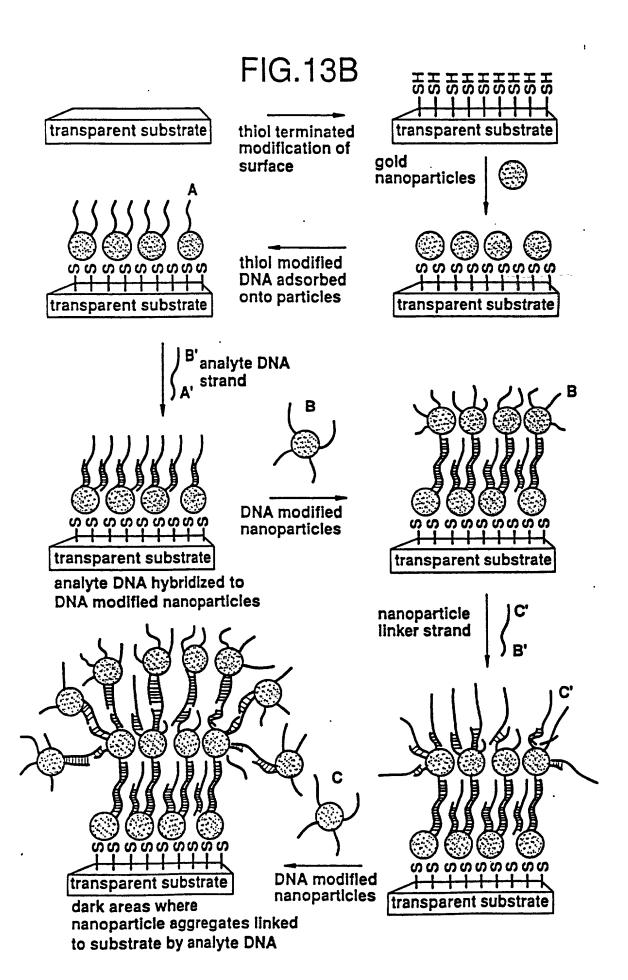


FIG.14A

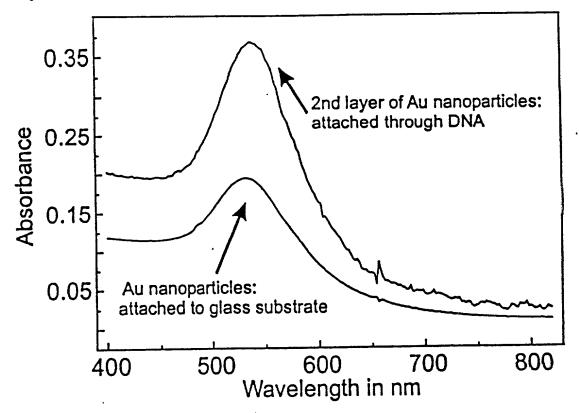


FIG.14B

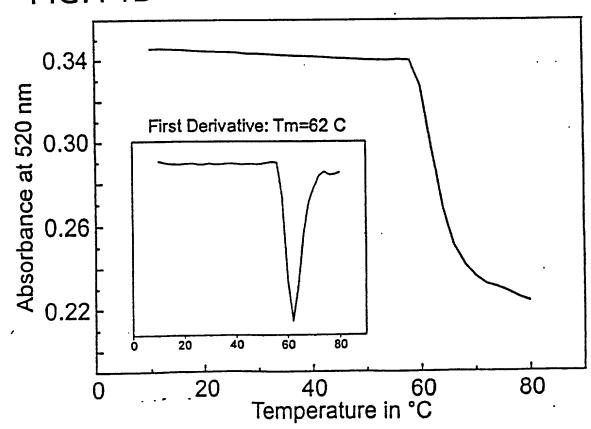


FIG15A
Probes with No Target SEQ ID NO:19 SEQ ID NO:20
S-ATG-CTC-AAC-TCT TAG-GAC-TTA-CGC-S
FIG15B
Half-Complementary Target SEQ ID NO:21
5' TAC-GAG-TTG-AGA-GAG-TGC-CCA-CAT 3'
S-ATG-CTC-AAC-TCT TAG-GAC-TTA-CGC-S
FIG15C
Complementary Target Tm=53.5°C
SEQ ID NO:22 5' TAC-GAG-TTG-AGA-ATC-CTG-AAT-GCG 3'
S-ATG-CTC-AAC-TCT TAG-GAC-TTA-CGC-S
FIG15D 2
ONE Base-Pair Mismatch at Probe Head Tm=50.4°C
5 SEQ ID NO:23
5' TAC-GAG-TTG-AGA-ATC-CTG-AAT-GC <u>T</u> 3' S-ATG-CTC-AAC-TCT TAG-GAC-TTA-CGC-S
<u>1</u> <u>2</u>
FIG15E Tm=46.2°C
ONE Base-Pair Mismatch at Probe Tail
6 SEQ ID NO:24 5' TAC-GAG-TTG-AGA-CTC-CTG-AAT-GCG 3'
S-ATG-CTC-AAC-TCT TAG-GAC-TTA-CGC-S 1 2
FIG15F ONE Base Deletion Tm=51.6°C
5' TAC-GAG-TTG-AGA-ATC-CTG-AAT-GC□3'
FIG15G
ONE Base-Pair Insertion 8 Tm=50.2°C SEQ ID NO:26
5' TAC-GAG-TTG-AGA-CAT-CCT-GAA-TGC-G 3'
S-ATG-CTC-AAC-TCT TA-GGA-CTT-ACG-C-S 1 2

FIG. 16A 24 Base Template

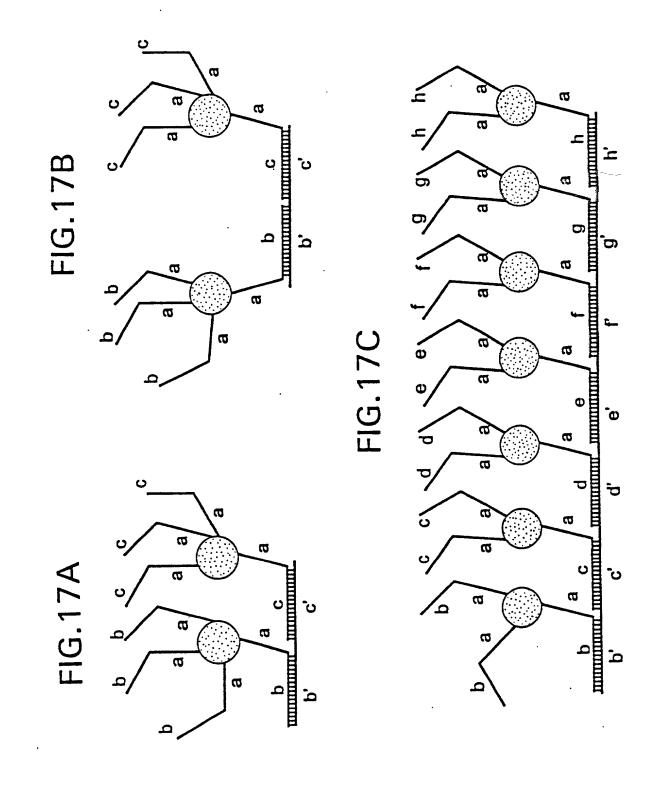
5' TAC-GAG-TTG-AGA-ATC-CTG-AAT-GCG 3'

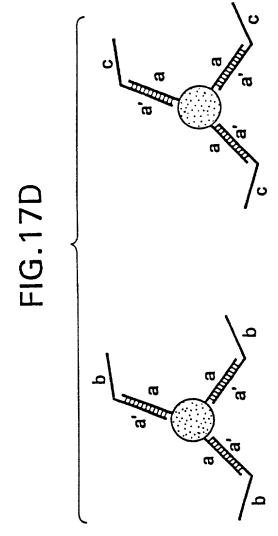
S-ATG-CTC-AAC-TCT TAG-GAC-TTA-CGC-S \to \frac{1}{2}

48 Base Template with Complementary 24 Base Filler FIG. 16B

5' TAC-GAG-TTG-AGA-CCG-TTA-AGA-CGA-GGC-AAT-CAT-GCA-ATC-CTG-AAT-GCG 3' S-ATG-CTC-AAC-TCT GGC-AAT-TCT-GCT-CCG-TTA-GTA-CGT TAG-GAC-TTA-CGC-S

72 Base Template with Complementary 48 Base Filler FIG. 16C





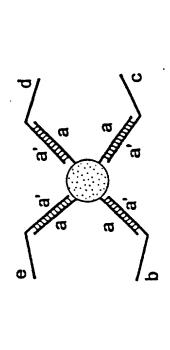
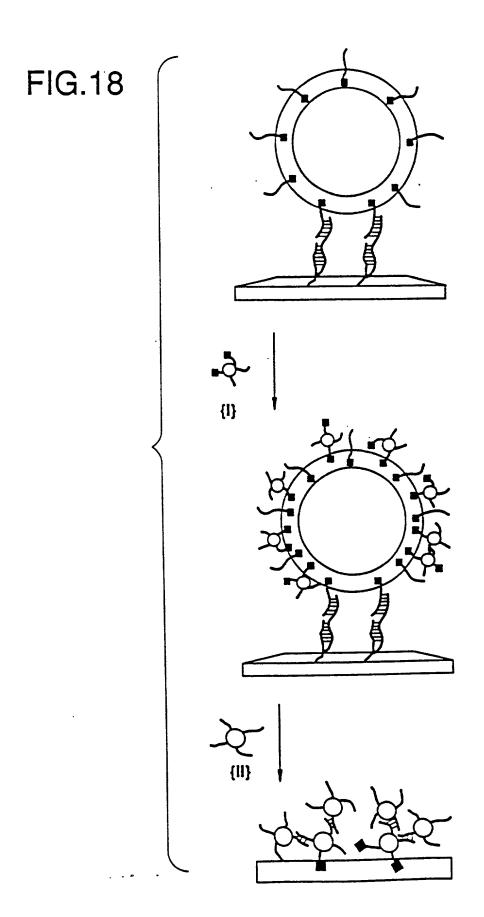
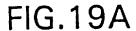


FIG.17E





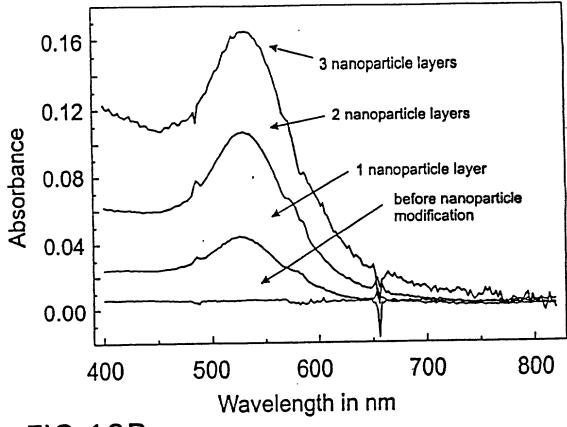
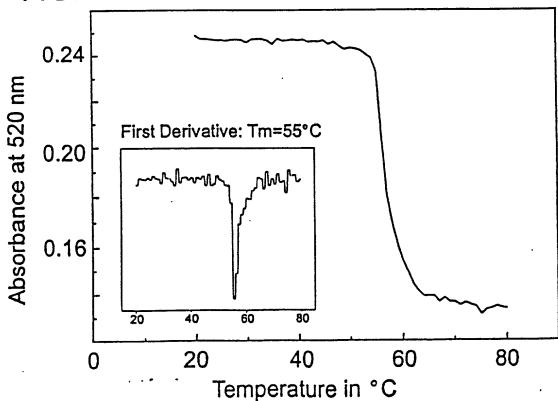
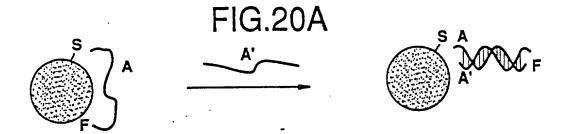
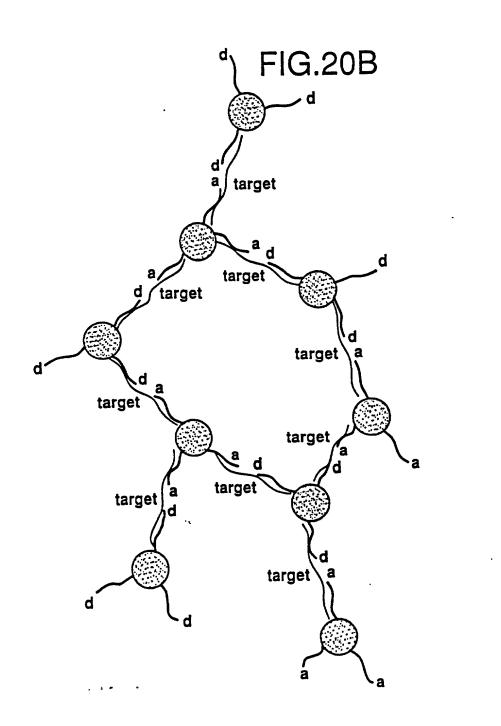


FIG.19B







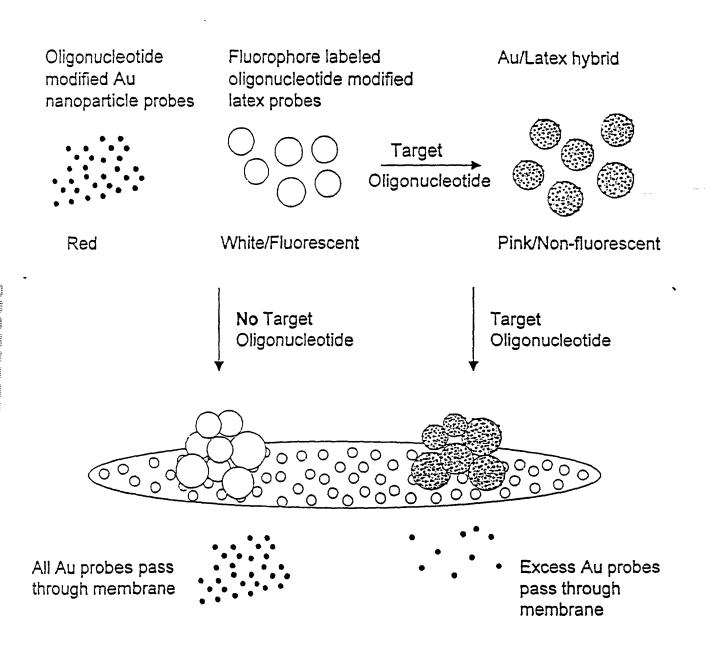
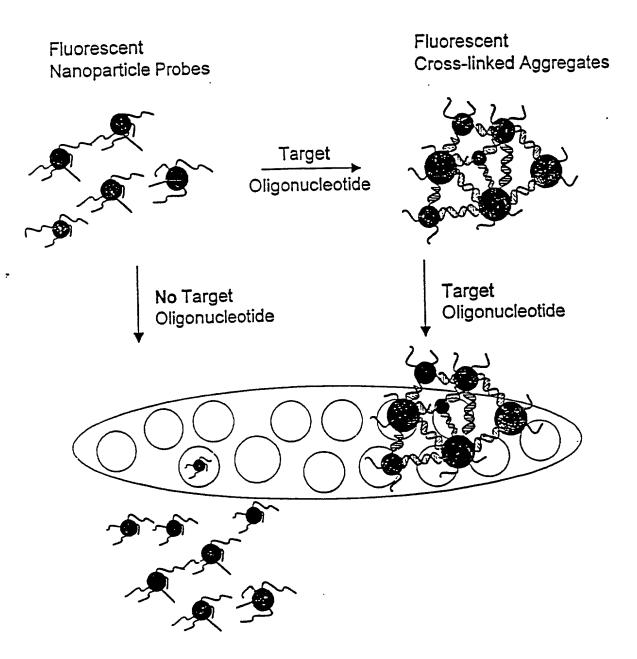


FIGURE 21



The fluorescent nanoparticle probes pass through the membrane

The fluorescent cross-linked aggregates are retained by the membrane

Anthrax PCR Product

5'G GCG GAT GAG TCA GTA GTT AAG GAG GCT CAT AGA GAA GTA ATT AAT 3'C CGC CTA CTC AGT CAT CAA TTC CTC CGA ĞŢA TCT CTT CAT TAA TTA

TCG TCA ACA <u>GAG GGA TTA TTG TTA AAT ATT GAT AAG GAT</u> ATA AGA AAA AGC AGT TGT CTC CCT AAT AAC AAT TTA TAA CTA TTC CTA TAT TCT TTT

ATA TTA TCC AGG GTT ATA TTG TAG AAA TTG AAG ATA CTG AAG GGC TT 3' TAT AAT AGG TCC CAA TAT, AAC ATC TTT AAC TTC TAT GAC TTC CCG AA 5'

141 mer Anthrax PCR product [SEQ ID 10:36]

3' CTC CCT AAT AAC AAT

[SEQ 10 NO:31]

3' TTA TAA CTA TTC CTA

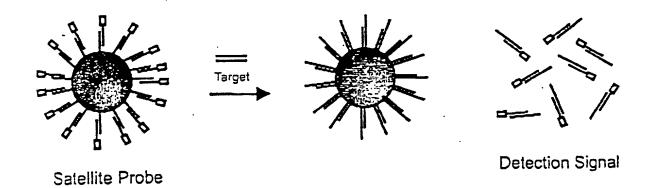
[SEQ ID NO: 38]

Oligonucleotide-Nanoparticle Probes

Blocker Oligonucleotides

3'C CGC CTA CTC AGT CAT CAA TTC CTC CGA GT [SEQ 1D NO:39]
3'A TCT CTT CAT TAA TTA AGC AGT TGT [SEQ 1D NO:40]
3'TAT TCT TTT TAT AAT AGG TCC CAA TAT [SEQ 1D NO:41]
3'AAC ATC TTT AAC TTC TAT GAC TTC CCG AA [SEQ 1D NO:42]

FIGURE 23



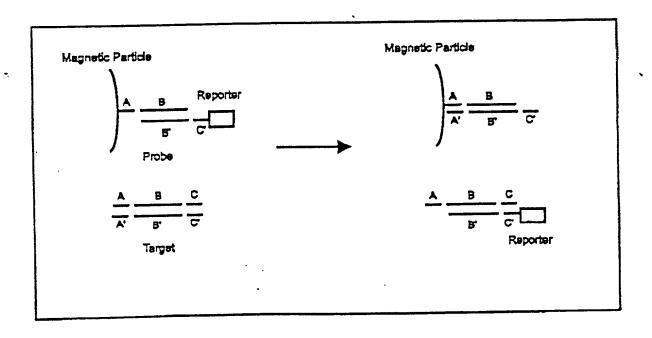
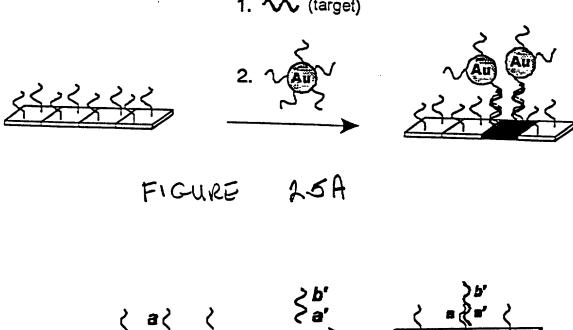
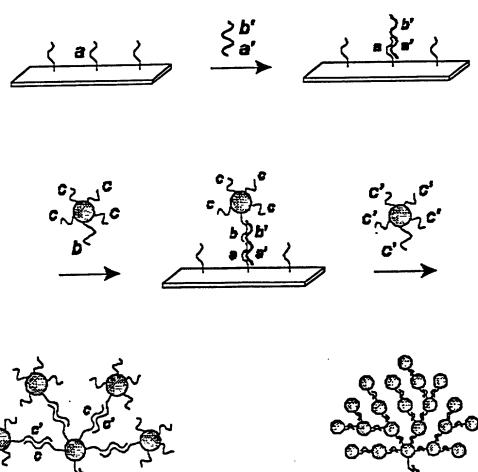


FIGURE 24

1. (target)





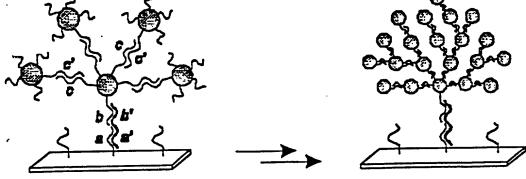


FIGURE 25 B

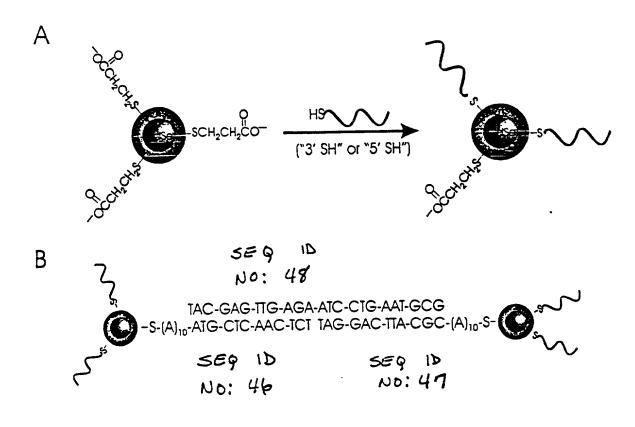
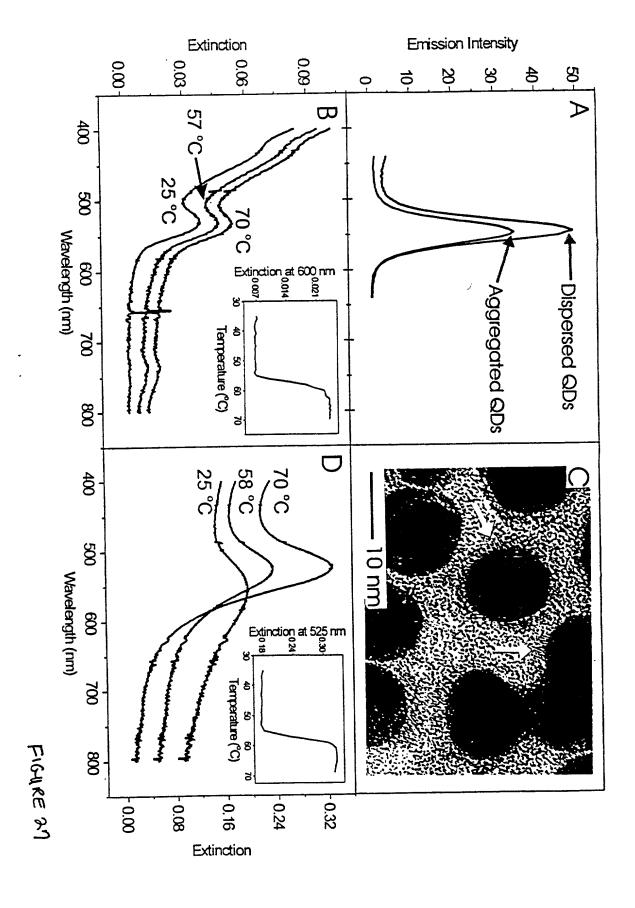
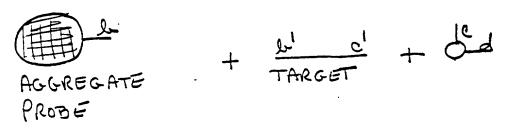
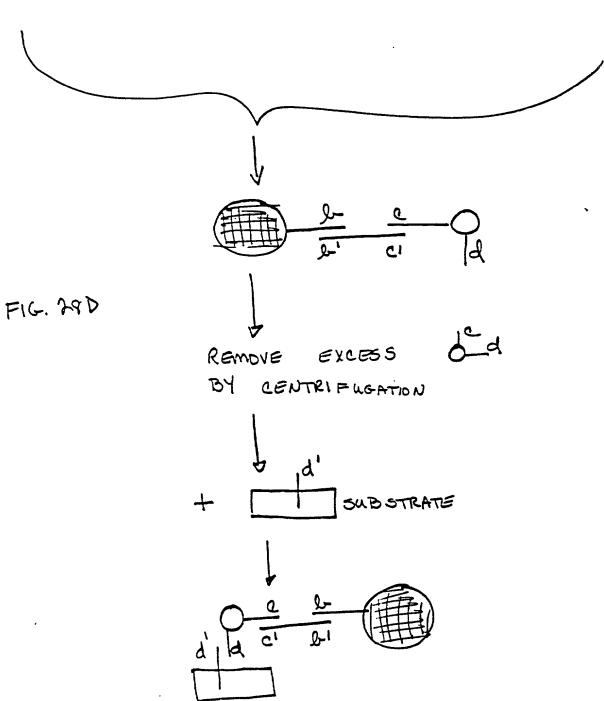


FIGURE 26





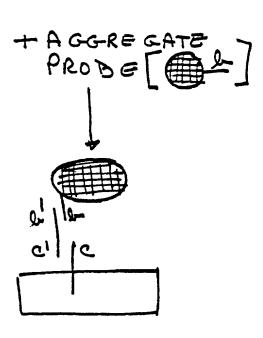


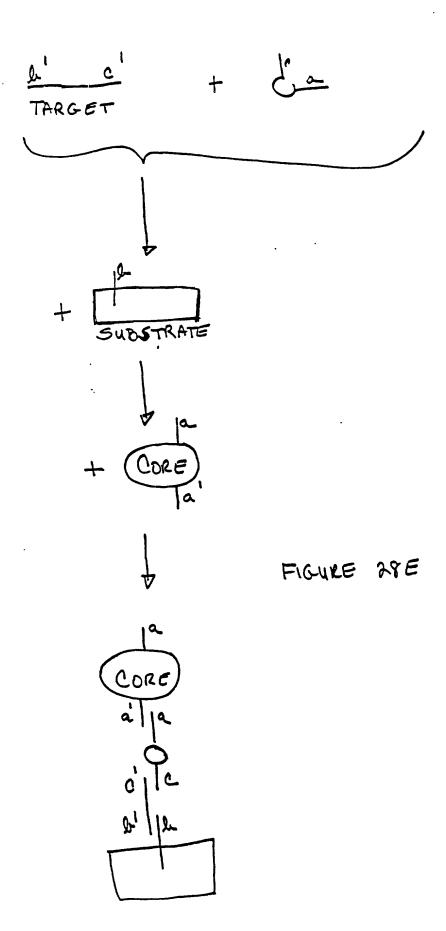
$$\frac{a - a}{a} + \frac{a - a}{a} - \frac{a}{a} \longrightarrow 0$$

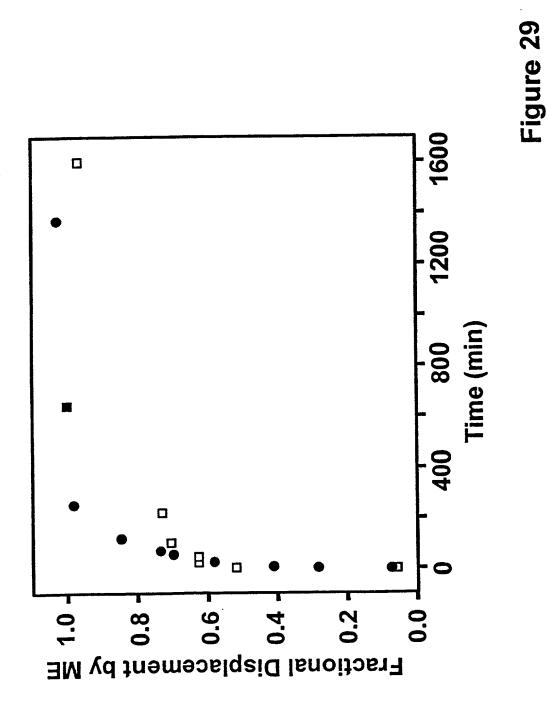
FIGURE 28A

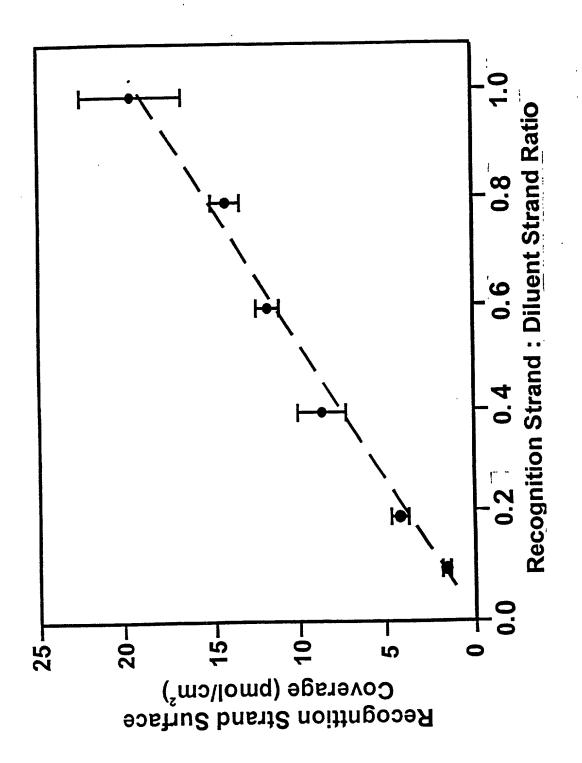
FIGURE B& B

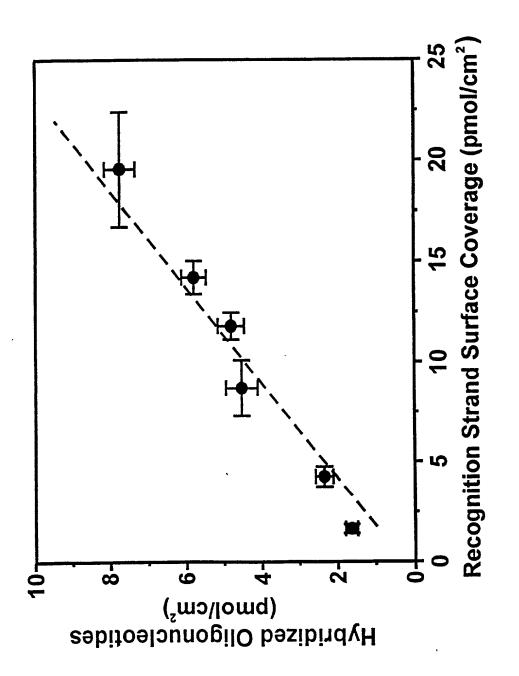
FIG. 28C











[SEQ ID NO:56]

5' GGA TTA TTG TTA---AAT ATT GAT AAG GAT 3'

CCT ANT AAC AAT TTA TAA CTA TTC CTA

[SEQ ID NO:57] [SEQ ID NO:58]

N = A (complementary), G,C,T (mismatched)

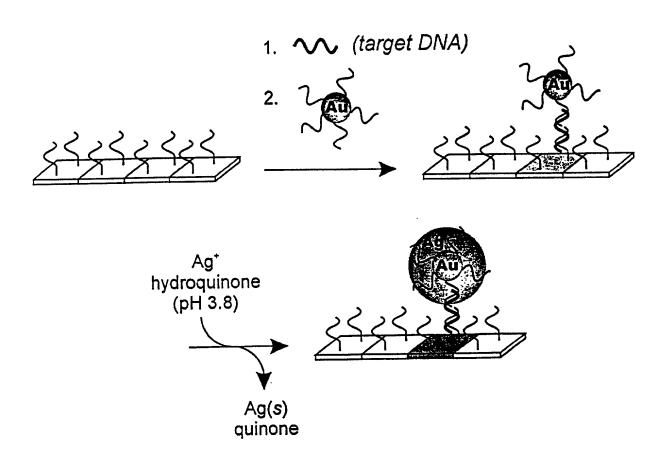


Figure 32

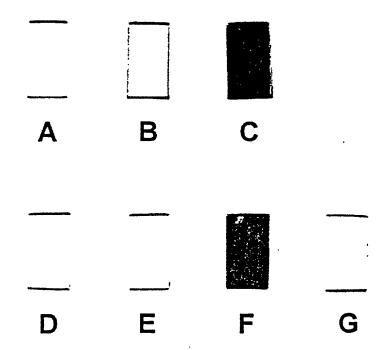
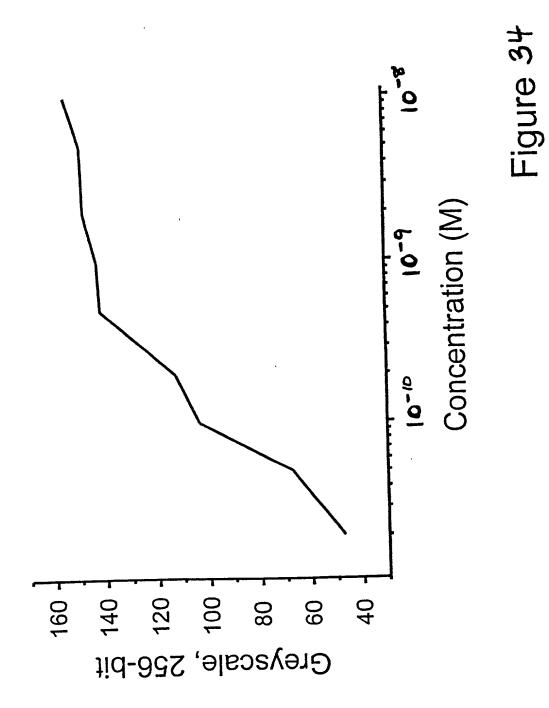


Figure 33



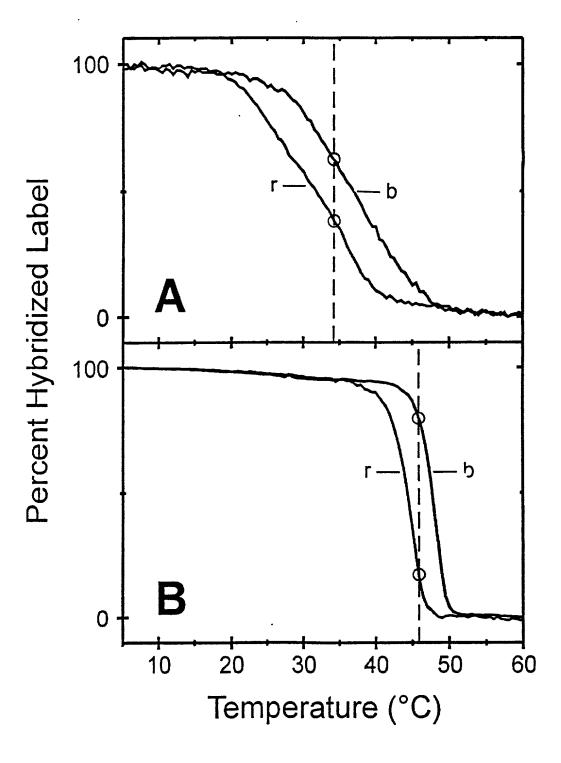


Figure 35

FIG. 364

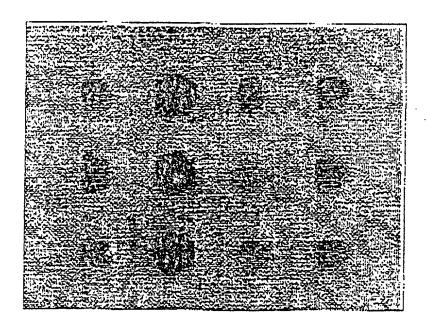
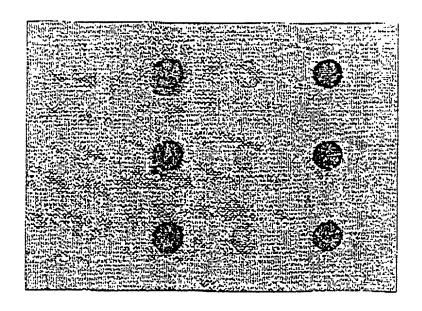


Fig. 368



C A T G

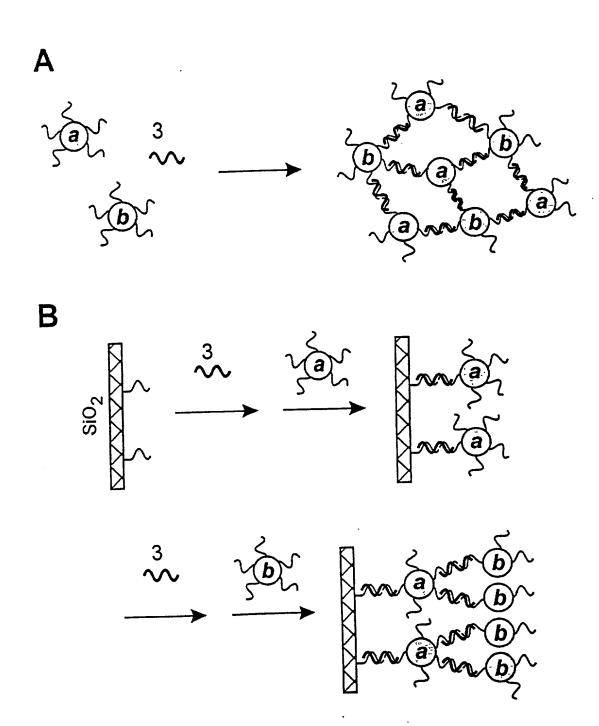


Figure 37

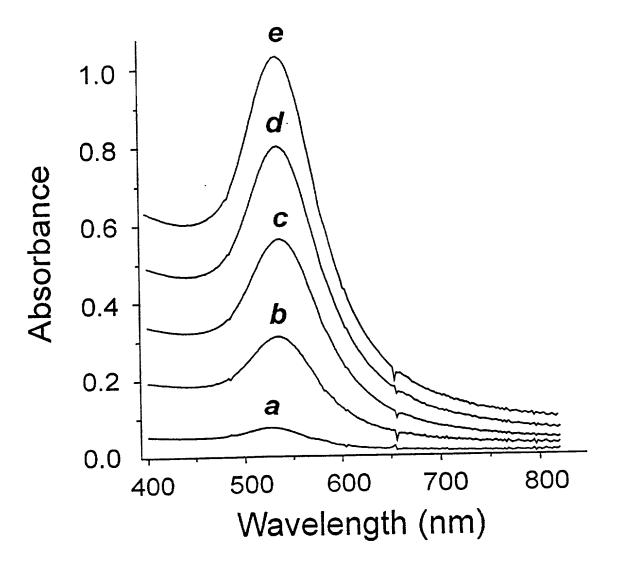


Figure 38A

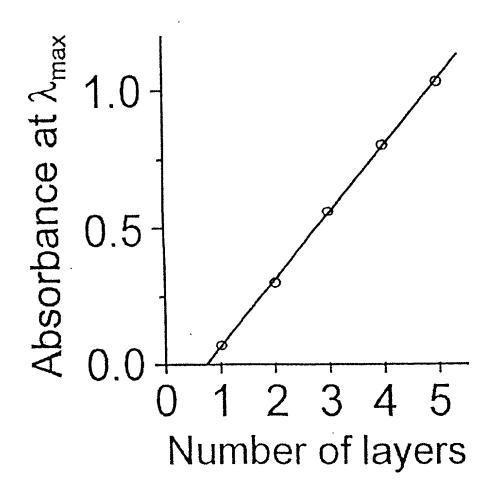


Figure 38B

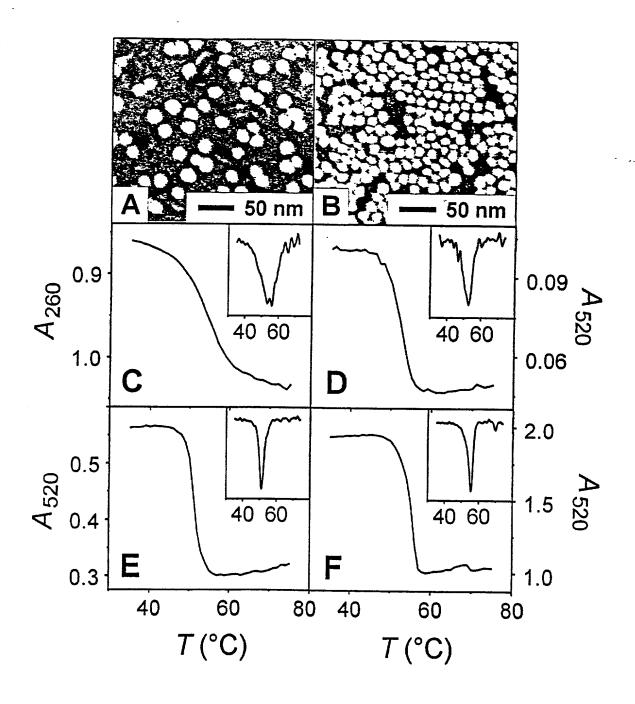
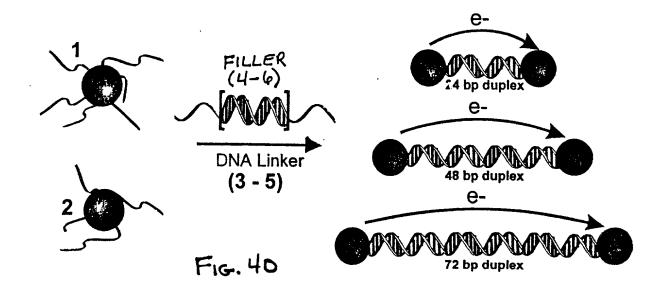


Figure 39



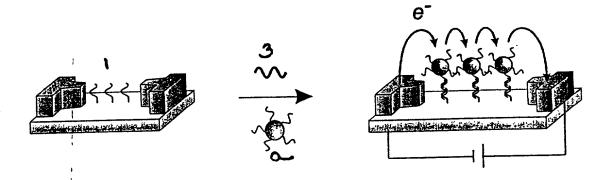


FIG. 41